

## Lecture 9

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(The odd numbered lectures are given by D. Ramakrishnan, and the even ones by R. Tanner.)

### 0.1 Comparison of $m$ with $m_{BH}$ and $m_{MM}$

Let  $m$  denote the slope at  $s = 0$  obtained in Lecture 7. Put

$$(6.1a) \quad m_{BH} := \frac{k_1 V}{k_2 + k_3},$$

which is the slope in the Briggs-Haldane model, and

$$(6.1b) \quad m_{MM} := \frac{k_1 V}{k_2},$$

the slope coming from the Michaelis-Menton model.

**Proposition** *We have*

$$m_{BH} \leq m \leq m_{MM}.$$

### 0.2 A quadratic approximation for small $s$

We explicated the linear term of our method in the previous lectures and compared it to the Briggs-Haldane and Michaelis-Menton approaches. Our approach can go much further and give an approximation of any order desired (for small  $s$ ). In the next section we will in fact give an exact infinite series expansion, from which approximations of any order can be deduced. Now let us delineate the quadratic case. We will preserve the earlier notations involving  $v_n, c_n$ , etc., denoting the values of  $v, c$ , etc., at the  $n$ -th stage of the recursion. Let us write

$$(6.2) \quad v_n = m_n s + q_n s^2 + O(s^3),$$

where  $m_n$  is the slope at the  $n$ -th stage, and  $q_n$  the coefficient controlling the quadratic term. As usual,  $O(s^r)$  denotes, for any  $r > 0$ , a sum of terms of order at least  $s^r$ . It follows by differentiation that

$$(6.3) \quad [v]'_n = m_n s' + 2q_n s s' + s' O(s^2) + O(s^3).$$

Since  $v = k_3 c$ , we get from (6.2),

$$(6.4a) \quad k_2 c_n = \frac{k_2}{k_3} m_n s + \frac{k_2}{k_3} q_n s^2 + O(s^3),$$

and

$$(6.4b) \quad k_1 s c_n = \frac{k_1}{k_3} m_n s^2 + O(s^3).$$

These give, thanks to (1) and (5), at stage  $n$ ,

$$(6.5) \quad s' = \left( \frac{k_2}{k_3} m_n - \frac{k_1}{k_3} V \right) s + \frac{(k_1 m_n + k_2 q_n)}{k_3} s^2 + O(s^3).$$

In particular,  $s'$  is  $O(s)$ , and

$$s s' = \left( \frac{k_2}{k_3} m_n - \frac{k_1}{k_3} V \right) s^2 + O(s^3).$$

So (6.3) simplifies as

$$(6.6) \quad k_3 c'_n = m_n s' + \left( \frac{k_2}{k_3} m_n - \frac{k_1}{k_3} V \right) s^2 + O(s^3).$$

The recursion is defined in such a way that

$$(6.7) \quad v_{n+1} = -s'_n - c'_n,$$

where  $s_n$  denotes the value of  $s'$  at stage  $n$ . Thus, by (6.6),

$$(6.8) \quad v_{n+1} = - \left( 1 + \frac{m_n}{k_3} \right) s'_n - \left( \frac{k_2}{k_3} m_n - \frac{k_1}{k_3} V \right) s^2 + O(s^3).$$

Plugging in (6.5), we then get

$$(6.9) \quad v_{n+1} = - \left( 1 + \frac{m_n}{k_3} \right) \left( \frac{k_2}{k_3} m_n - \frac{k_1}{k_3} V \right) s - \left( 1 + \frac{m_n}{k_3} \right) \left( \frac{k_1 m_n + k_2 q_n}{k_3} \right) s^2 - \left( \frac{k_2}{k_3} m_n - \frac{k_1}{k_3} V \right) s^2 + O(s^3).$$

Since this is  $m_{n+1}s + q_{n+1}s^2 + O(s^3)$ , we obtain

$$(6.10) \quad q_{n+1} = - \left( 1 + \frac{m_n}{k_3} \right) \left( \frac{k_1 m_n + k_2 q_n}{k_3} \right).$$

Now we let  $n$  go to infinity, and obtain, for  $q = \lim_{n \rightarrow \infty} q_n$ , the relation

$$q = - \left( 1 + \frac{m}{k_3} \right) \left( \frac{k_1 m + k_2 q}{k_3} \right).$$

This yields

$$(6.11) \quad q = - \frac{\left( 1 + \frac{m}{k_3} \right) \frac{k_1 m}{k_3}}{1 + \left( 1 + \frac{m}{k_3} \right) \frac{k_2}{k_3}}.$$

Note that this is independent of  $e_0$ .

Putting  $v = \lim_{n \rightarrow \infty} v_n$ , this gives the expression

$$(6.12) \quad v = ms - \left( \frac{(k_3 + m)k_1 m}{k_3^2 + (k_3 + m)k_2} \right) s^2 + O(s^3),$$

where  $m$  is given by (5.12b). Consequently,

$$(6.13) \quad \frac{dv}{ds} = m - \left( \frac{2(k_3 + m)k_1 m}{k_3^2 + (k_3 + m)k_2} \right) s + O(s^2).$$

If  $s_0$  is small, then so is  $S_p$ , in which case this quadratic approximation has some validity near it, allowing to deduce that  $S_p$  is close to  $\frac{k_3^2 + (k_3 + m)k_2}{2(k_3 + m)k_1}$ .